(12) INTERNATIONAL APPLICATION PUBLISHED UNDER THE PATENT COOPERATION TREATY (PCT)

(19) World Intellectual Property Organization International Bureau





(43) International Publication Date 16 May 2002 (16.05.2002)

PCT

(10) International Publication Number WO 02/38040 A2

(51) International Patent Classification7:

A61B 1/00

(21) International Application Number: PCT/US01/49704

(22) International Filing Date:

9 November 2001 (09.11.2001)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data: 09/709,162

10 November 2000 (10.11.2000) U

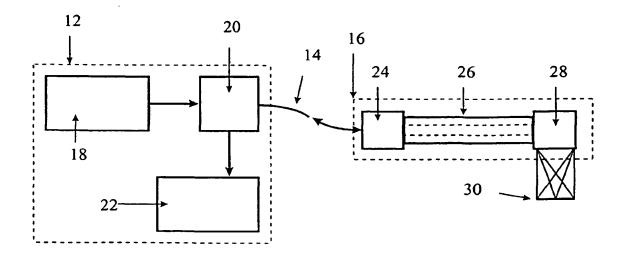
- (71) Applicant (for all designated States except US): THE GENERAL HOSPITAL CORPORATION [US/US]; 55 Fruit Street, Boston, MA 02114 (US).
- (72) Inventors; and
- (75) Inventors/Applicants (for US only): TEARNEY, Guillermo, J. [US/US]; 118 Kinnaird Steet Number 3, Cambridge, MA 02139 (US). BOUMA, Brett, Eugene [US/US]; 12 Monmouth Street, Quincy, MA 02171 (US).

SHISKOV, Milen, Stefanov [BG/US]; 122 Bowdoin Street, Apt. 2, Boston, MA 02108 (US). ROSEN, Jonathan, Jay [US/US]; 12 Surrey Road, Newton, MA 02458 (US).

- (74) Agent: BERNSTEIN, Jason, A.; Bernstein & Associates, P.C., Suite 495, Embassy Row 400, 6600 Peachtree Dunwoody Road, N.E., Atlanta, GA 30328-1649 (US).
- (81) Designated States (national): AE, AG, AL, AM, AT, AT (utility model), AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, CZ (utility model), DE, DE (utility model), DK, DK (utility model), DM, DZ, EC, EE, EE (utility model), ES, FI, FI (utility model), GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SK (utility model), SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE,

[Continued on next page]

(54) Title: SPECTRALLY ENCODED MINIATURE ENDOSCOPIC IMAGING PROBE



(57) Abstract: A spectrally encoded endoscopic probe having high resolution and small diameter comprising at least one flexible optical fiber; an energy source; a grating through which said energy is transmitted such that the energy spectrum is dispersed; a lens for focusing the dispersed energy spectrum onto a sample such that the impingement spot for each wavelength is a separate position on the sample, the wavelength spectrum defining a wavelength encoded axis; means for mechanically scanning the sample with focused energy in a direction perpendicular to the wavelength encoded axis; a means for receiving energy reflected from the sample; and, a means for detecting the received reflected energy. The probe grating and lens delivers a beam of multispectral light having spectral components extending in one dimension across a target region and which is moved to scan in another direction. The reflected spectrum is measured to provide two dimensional imaging of the region.

Ø 07/38040 C

IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

For two-letter codes and other abbreviations, refer to the "Guid- ance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

Published:

 without international search report and to be republished upon receipt of that report

BNSDOCID: <WO____0238040A2_I_>

5

SPECTRALLY ENCODED MINIATURE ENDOSCOPIC IMAGING PROBE

10

15

Field of the Invention

The present invention relates to the field of endoscopy in general and to an endoscope having a combination of high number of resolvable points using spectral encoding and contained in a diameter/space small enough to perform desired procedures.

20 Background of the Invention

Clinical use of endoscopic devices and probes has permitted physicians to view and diagnose target bodies, such as tumors, deposits, tears, thrombi, and the like. Unfortunately, there are still a number of disadvantages and limitations of using conventionally available endoscopes. The diameter of currently available probes limits their use to certain procedures and locations that can accommodate the large diameter of the endoscope. Consequently, many procedures currently done surgically

PCT/US01/49704 WO 02/38040

could be done endoscopically, if a small enough probe was available with a sufficient number of resolvable points.

Current endoscopic procedures generally require administration of anesthesia and surgical training for insertion. Certain procedures cannot currently be done endoscopically because of the diameter of existing probes. Probe size is proportional to the likelihood of tissue damage occurring during the procedure. Another problem associated with current probes is the occurrence of adverse reactions, such as in fetoscopy, where the risk to the fetus is high. Neural imaging carries with it the possibility of brain damage. Spinal canal and brain ventricular imaging have complications of spinal fluid leakage and headaches which are more frequent and severe with larger diameter probes. Catheterization of the pancreatic duct is also problematic due to the probe size and resultant complications which include acute pancreatitis. Also, the size of the incision necessary to insert current probes results in longer healing time and more prominent scarring.

Present day miniature endoscopes are composed of fiber-optic imaging bundles. Currently, the clinical use of small diameter endoscopes is limited by poor resolution. Available miniature endoscopes have diameters in the range of from about 0.35 mm to about 1.0 mm. Since optical fibers are of finite diameter, only a limited number of fibers can be incorporated into one imaging bundle, resulting in a limited number of resolvable elements. For example, for a 1 mm fiber optic imaging bundle with an individual fiber diameter of 10 μm , the total number of resolvable points is 9000 with 100 resolvable points across the field of view. In addition, the fill factor is about 85% resulting dead space from the cladding material and causing the image to have a pixelated or "honeycomb" appearance. These two technical problems have severely limited the clinical use of currently available sub-millimeter diameter imaging probes. In order to achieve a higher number of resolvable elements with such probes, larger diameters must be used, which obviate their use in smaller spaces and eliminate certain procedures from being done endoscopically. If one uses a currently available probe in the sub-millimeter diameter range, the number of resolvable points obtainable drops below clinically useful levels. Present endoscopes have a light 30 transmission efficiency of up to about 50%.

A further disadvantage of fiber bundles is that crosstalk occurs, reducing the signal to noise level. Moreover, as fiber length increases, light transmission efficiency

5

10

15

20

decreases. Also with current fiber optic endoscopes, coupling illumination light into the fiber optic imaging bundle is difficult. As a result, miniature endoscopes need two separate fiber bundles, one for illumination and one for detection of the image. The need for distinct illumination and detection bundles increases (at least doubles) the overall endoscope diameter. It would therefore be desirable to have a long length endoscope probe that retains sufficient light transmission efficiency to provide a clinically useful image and information.

Another disadvantage of optical fiber bundles is that individual fibers may be broken or have defects at their faces, resulting in "dead" pixels. The use of one fiber would greatly minimize the presence of dead pixels.

Thus, it would be desirable to have a probe that provided a satisfactory number of resolvable elements in a space/diameter below a certain size that would enable procedures to be done currently not achievable by currently available endoscopes. It would be desirable to have a probe in the sub-millimeter diameter range that had optics that would improve the number of resolvable points, reduce deadspace/fill factor, and minimize risk of adverse consequences to the patient. Such a novel probe would enable procedures to be performable that currently cannot be attempted endoscopically, such as, but not limited to, otological, neural, pancreatic, and fetal surgical endoscopy. It would also be desirable to have a sub-millimeter endoscope that would allow for diagnosis as well as treatment in a single device.

Spectral encoding is a method that allows detection of a one-dimensional line of an image using a single optical fiber. Encoding the spatial information on the sample is accomplished by using a broad bandwidth source as the input to the endoscope. The source spectrum is dispersed by a grating and focused by a lens onto the sample. The spot for each wavelength is then focused at a separate position, x, on the sample. The reflectance as a function of transverse location is determined by measuring the reflected spectrum. The other dimension of the image can be obtained by mechanical scanning at a slower rate. The advantage of this mode of imaging is that the fast scanning needed to produce an image at or near video frame rates is performed externally to the probe, making the construction of small diameter probes feasible.

5

10

15

20

25

PCT/US01749704 WO 02/38040

Summary of the Invention

5

10

15

20

25

The present invention discloses a miniature imaging probe or endoscope that is capable of obtaining real-time images with up to or higher than about 100 times the number of resolvable points than a fiber-optic imaging bundle of the same diameter. Another way of describing the present invention is for the same number of resolvable pixels as commercially available imaging bundles, the present invention could have a diameter that is 10 times smaller. In addition, this instrument produces images that do not contain cladding artifacts. Also, this invention allows acquisition of depth (three-dimensional, or 3D) information from the sample. Finally, by acquiring multiple images of the sample, spectroscopic information relating to the chemical composition of the object may be obtained. These properties make this device an enabling technology for performing endoscopic or catheter based imaging in previously inaccessible locations within the body. Important specific applications include fetoscopy, pediatric endoscopy, coronary angioscopy, mini-laparoscopy, mammary ductoscopy, lacrimal ductoscopy, small joint visualization, and other medical and non-medical applications.

a cian Comini
Significant work related to this invention is described in copending application, Serial
Significant work related to this myodian to regular application, Serial
No. 60/076,041, filed February 26, 1998 (corresponding to regular application, Serial No. 60/076,041, filed February 26, 1998 (corresponding to regular application, Serial No.
and International Approximation
No No 26 1999) entitled Confocal Microscopy With
No, filed, need, need
the disclosures of which are incorporated have
described the use of special checkers
entirety. This prior disclosure described are entirety.
endoscopic confocal microscopy. The present invention description macroscopic
11 diameter endoscopes and obtain ingit-tooks.
encoding to create small diameter chaososper and applications of images. The present invention includes different embodiments and applications of images.
images. The present invention includes different small diameter probes.
spectral encoding for performing endoscopic imaging through small diameter probes.

Objects of the Invention

Accordingly, it is a principal object of the present invention to provide an endoscope having a combination of resolution above a certain level in a diameter or space below a certain size.

It is an object of the present invention to provide an endoscope having a combination of number of resolvable elements above about 10,000 in a space below about 1.0 mm².

It is another object of the present invention to provide an endoscope in the generally sub-millimeter diameter range that increases the number of resolvable elements.

It is also an object of the present invention to provide an endoscope that can be used in smaller spaces than currently available endoscopes.

It is another object of the present invention to provide an endoscope that can be used in conjunction with a small diameter needle so as to avoid anesthesia and where the endoscopic procedure can be done, where appropriate, on an outpatient basis.

It is another object of this invention to provide an endoscope which is simpler to manufacture and is less likely of having individual pixel defects than current fiber optic bundles.

It is a further object of the present invention to provide an endoscope using spectral encoding and using a single fiber.

It is yet another object of the present invention to provide an endoscope capable of obtaining spectroscopic information from the sample.

It is a further object of the present invention to provide a third dimension of information (depth), above the typical two dimensions of data obtainable from conventional endoscopes.

It is yet another object of the present invention to provide higher sensitivity and higher signal to noise ratio images for clearer visualization through turbid media.

It is still another object of the present invention to provide an endoscope that can reduce the likelihood of tissue damage and other adverse consequences.

It is another object of the present invention to provide an endoscope that allows for diagnosis and treatment in a single procedure.

It is yet a further object of the present invention to provide an endoscope having no fill factor problem.

-5

10

15

PCT/US01/49704 WO 02/38040

It is yet another object of the present invention to provide an endoscope with no "dead" pixels.

Other objects, features and advantages of the present invention will become apparent upon reading the following detailed description of embodiments of the invention, when taken in conjunction with the appended claims.

Brief Description of the Drawings

5

The invention is illustrated in the drawings in which like reference characters designate the same or similar parts throughout the figures of which:

Fig. 1 is a schematic view of a spectrally encoded miniature camera according to a preferred embodiment of the present invention.

Figs. 2A-F are schematic views of different possible configurations of the distal probe of the camera of Fig. 1.

Figs. 3A-C are schematic views of the image formation geometry for linear, sector and circular scans, respectively.

Fig. 4 is a schematic view of a preferred embodiment of the detection system of the camera of Fig. 1.

Fig. 5 is a schematic view of an alternative embodiment of the present invention showing a color spectrally encoded miniature camera.

Fig. 6 is a schematic view of an alternative embodiment of the present invention showing a multifiber array incorporating a plurality of grating lenses.

Fig. 7 is a schematic view of an alternative embodiment of the present invention showing an optical fiber having spaced lenses along its length.

Description of the Preferred Embodiments

The present invention provides in general an endoscopic probe having a small diameter and a high number of resolvable elements. For the purposes of the present disclosure, a probe diameter in the sub-millimeter range is most desirable; however, it is to be understood that diameters of or greater than 1 mm can be used with the

present invention. The advantages of the resolution of the present invention certainly carry to probes with greater than 1 mm diameter. Thus, the present invention provides a combination of number of resolvable points above a certain level in a diameter or space below a certain level.

Fig. 1 shows an endoscope 10 of a preferred embodiment consisting of main body 12 connected through a hybrid (optical and electrical) cable 14 to a probe 16. The cable 14 can instead be a discontinuity conducted across a gap (for example, optical, electrical or electromagnetic relay), discussed hereinbelow with respect to an alternative embodiment using a self-contained probe with a transmitter to a remote receiver/detector. The main body 12 incorporates a broadband source 18 that sends the illuminating light to the beam-redirecting element 20. In different designs this beam-redirecting element 20 can be a beam splitter (simple, inexpensive, but poor light efficiency), a polarizing splitter, an optical circulator or other device known to those skilled in the art. This beam-redirecting element 20 sends out the light from the source 18 to the probe 16 and redirects the returning light from the probe 16 to the detection and display system 22. The detection system is preferably associated with a computer with a microprocessor (not shown).

The probe 16 has three major components. The proximal end 24 generates the onedimensional scanning. In a preferred embodiment the scanning is done mechanically. A flexible, semi-flexible or rigid tube 26 connects the proximal end 24 and the distal end 28 of the imaging probe 16. An optical fiber conveys the optical signals and a hollow flexible, semi-flexible, or rigid cable conveys one-dimensional scanning of the sample 30.

The cable 14 is preferably a single mode optical fiber. Another embodiment is a double-clad fiber where the illumination is through the central core and the detection is through the central core and outer cladding. Alternatively, the cable 14 can be a co-axial or side-by-side pair of fibers, with one fiber being for illumination by the source and the other fiber being for collection of light reflected from the sample 30. In a device where a multimode fiber system is used, there may be increased sensitivity and decreased speckling. Speckle artifacts may also be reduced by introducing mode or phase modulation of the fiber. Mode or phase modulation may be performed by rapidly moving the optical fiber, or insertion of an optical element in the fiber path capable of rapidly changing modes or phase of the probe light. This embodiment

5

10

15

20

25

PCT/US01/49704 WO 02/38040

includes the detection mechanism within the probe and a RF transducer to relay image information remotely to a RF detector).

Sources

5

10

20

25

The light source 18 can be any broadband source capable of performing highresolution imaging using the spectral encoding method. Examples of sources include, but are not limited to, light-emitting diodes, super-luminescent diodes, rare-earth doped fibers, solid-state mode-locked lasers, spectrally broadened laser sources, wavelength tunable light sources, monochromatic light, polychromatic light, and the like. The source 18 does not need to illuminate all wavelengths simultaneously. It can emit a monochromatic radiation whose wavelength is scanned with time. This allows fluorescence illumination to be done. It also does not require Fourier transform to be done. It is to be understood that other sources of energy can be used, including, but not limited to, infrared, ultraviolet, ultrasonic, low or high energy radiation (for example, x-ray, alpha, beta, gamma, and the like), other electromagnetic radiation, combinations of all of the foregoing and the like. For higher energy radiation, certain of the components of the present invention may need to be adapted for greater 15 shielding or other functional characteristics.

Distal Probe Design

The distal probe 16 design of the present invention will now be discussed. Fig. 2 shows several possible configurations of the imaging head 28. The optics of the head 28 is designed to produce linear, spectrally encoded illumination and to collect the reflected light and transmit it back to the detection system 22. The light from the source 18 is delivered by the fiber 14 to the head 16 and focused by an objective 32 onto the sample 30. In a preferred embodiment the objective 32 is a lens, for example, but not by way of limitation, a GRIN (gradient index) lens, as is known to those skilled in the art. Other possible lens elements include, but are not limited to aspherical lenses, planoconcave, biconcave, concaveconvex or multi-element lens assemblies.

Immediately after (or before) the objective 32, a grating 34 is used to disperse the source spectrum. In a preferred embodiment, shown in Fig. 2A, the grating 34 is a holographic grating. Alternatively, blazed or binary gratings or a grism (grating prism or carpenter's prism) can be used. Holographic gratings, however, are believed to be

8

better and have higher efficiency than blazed gratings for the intended use. Fiber gratings may also be used. The spot for each wavelength is focused at a separate position on the sample. The reflectance as a function of transverse location is determined by measuring the reflected spectrum. The head 16 also provides one-dimensional mechanical scanning orthogonal (or other angle) to the spectrally encoded axis. Spectral dispersion in one dimension while scanning the other dimension provides two-dimensional illumination of the sample.

Figs 2B-F illustrate alternative embodiments of the probe 16 design of Fig. 2A. On the Figures the elements are: optical fiber 14; imaging head 16; objective 32; diffracting grating 34 (Fig. 2A and 2D show a transmission grating, Figs. 2B, C and E show a reflecting grating 36, and Fig. 2F shows a fiber 38); a beam stop 40; a spectrally encoded imaging line 42; a mirror 44; a beam splitter 46; a polarizer 48; a polarizing beam splitting cube 50; and a 1/4 wavelength plate.

Image formation will now be discussed. The use of the dispersing element 34 and the focusing of the spectrum on the sample 30 to be imaged produces a one-dimensional scan. In order to obtain a two-dimensional image, one must perform a transverse scan in the conjugate direction. This can be implemented in many different embodiments, but all include a means of moving the spectrally encoded scan line. In a preferred embodiment the movement is achieved mechanically. Several possible methods for one-dimensional mechanical scanning are shown in Figs. 3 A-D. It is possible to use a piezoelectric transducer or a torque-transducing device known to those skilled in the art to achieve movement. The probe body 12 is associated with a transparent cap 60. A flexible cable 62 carries the fiber 14. The optical head 28 and the spectrally encoded imaging line 64 are shown. The two-dimensional imaging region is shown at 66. Fig. 3A shows a linear scan; Fig. 3B shows a sector scan; Fig. 3C shows a circular scan; and, Fig. 3D shows a forward circular scan. This may be performed by use of a rotating or linearly translating cable where the motion is produced at the proximal end of the endoscope. One can either rotate the optical head or push it linearly forth and back, moving the imaging line on the sample and scanning the image. The fiber may also be moved as opposed to the distal optics to give the alternate scan.

The proximal probe end design and coupling mechanisms will now be discussed. Referring back to Fig. 3, the proximal end 24 of the probe 16 provides the mechanical scanning necessary for obtaining a two dimensional image. In addition, this end

5

10

15

20

25

couples the light coming from the source 18 to the distal end 28 of the probe 12 and the optical signal coming back from the sample 30. Linear (axial or radial), sector, or circumferential scanning patterns are possible depending on the application. For example, in narrow vessel imaging, such as angioscopy, circumferential scanning is necessary to image the total surface of the vessel. In this case a special rotating junction must be incorporated into the proximal end 24 of the probe 16 to connect optically a stationary and a rotating fiber 14. For sector and linear scanning, the optical fiber14 within the probe 16 may be directly fused to the beam-redirecting element 20. The preferred scanning embodiment is sector or circumferential scanning due to the availability of small diameter cables already constructed for this purpose.

Resolution

5

10

The number of wavelengths or points that may be resolved is determined by:

$$\frac{\lambda}{\partial \lambda} = mN, \tag{1}$$

where λ is the center wavelength, $\partial \lambda$ is the bandwidth of the spectrum, N is the number of lines in the grating illuminated by the polychromatic input beam, and m is the diffraction order. If the total bandwidth of the source is $\Delta \lambda$, the number of resolvable points, n is defined by:

$$n = \frac{\Delta \lambda}{\partial \lambda} \,. \tag{2}$$

For an input source with a center wavelength of 1000 nm, a bandwidth of 200 nm, an input spot diameter of 1 mm, a diffraction grating of 1200 lines/mm and a diffraction order of 1, n = 240 points may be resolved by the spectrally encoded system. Moreover, if the grating is at an angle (θ) with respect to the incoming light, the number of resolvable points scales with 1/cos(θ). In the above example, for an incident angle of 65°, the number of resolvable points would be approximately n = 570. If the perpendicular direction was also scanned to give 570 points of resolution, the total number of resolvable points would be approximately 320,000. This is compared to 10,000 resolvable points (n = about 80) found in state of the art single

mode fiber bundles with a diameter of 1 mm. Thus, the present invention can provide approximately 16 times better resolution than conventional fiber optic probes.

The parameters used in this example may be found in common, inexpensive optical components. The number of points may be increased by simply increasing the input angle or the bandwidth of the source 18. Increasing the spot diameter increases the resultant probe diameter. Increasing the bandwidth of the source 18 could be accomplished by using a broader bandwidth superluminescent diode, a broad bandwidth LED, a rare earth doped fiber superfluorescent source, a solid state modelocked laser, a continuum source or the like.

Practitioners in endoscopic procedures would generally agree that a minimum number of resolvable elements of about 256 x 256 = 65,536 is needed to do meaningful diagnostic procedures. This is based on the fact that this is approximately the current accepted standard used in laparoscopic procedures. Current high end endoscopes have an upper end of about 10,000 elements in about a 1 mm diameter probe. Currently also, there are several fiber bundles that have 30,000 elements in about a 1 mm probe, but they are not yet in clinical use due to technical limitations.

The device of the present invention provides, in a preferred embodiment, a probe having a diameter of about 1.0 mm with a number of resolvable points in the range, in its broadest aspect, of from about 10,000 to about 1,000,000 resolvable points, more preferably, of from about 300,000 to about 1,000,000 resolvable points, more preferably of from about 150,000 to about 300,000 resolvable points, and still more preferably of from about 100,000 to about 150,000 resolvable points. The number of resolvable points roughly scales with diameter. It is to be understood by those skilled in the art that other diameters can be used with correspondingly greater or lesser numbers of resolvable points.

Table 1 provides a comparison of the number of resolvable elements using a spectrally encoded endoscope ("SEE") of the present invention compared to that using conventional fiber optic bundles. For the SEE, $\Delta\lambda$ (bandwidth) 250 nm, Λ (grating density) 1200 lines per mm.

5

10

15

20

PCT/US01/49704 WO 02/38040

TABLE 1

Diameter (mm)	Number of resolvable elements	
	Fiber Bundle	Spectral Encoded Endoscope
1.0	10000	160000
0.7	5000	80000
0.5	2500	40000
0.25	625	10000
0.23		

Energy budget

The intensity *I* of the returned light in a Michelson interferometer configuration of the camera is given by

$$I = \frac{1}{8} \frac{\gamma_1^2 \gamma_2 d^2 \rho \varphi}{b^2 N} I_0, \tag{3}$$

where I_0 is the source intensity, γ_1 and γ_2 are the diffracting efficiencies of the imaging and detecting grating, d is the beam diameter at the objective, ρ is the reflectivity of the surface to be imaged (spectral and location dependent), φ is a detector filling factor, and N is the number of the points to be detected.

The attenuation of the signal in $dB \ \epsilon$ is then given by

$$\varepsilon = 10 \log \frac{I}{I_0} = 10 \log \left[\frac{1}{8} \frac{\gamma_1^2 \gamma_2 d^2 \rho \varphi}{b^2 N} \right]$$
 (4)

In this formula different contributions can be considered as follows

$$\varepsilon = 10\log\frac{1}{8} + 10\log G + 10\log\Gamma + 10\log\rho + 10\log D,$$
 (5)

where $G = \gamma_1^2 \gamma_2$ is the diffracting efficiency factor, $\Gamma = \frac{d^2}{b^2}$ is the numerical aperture factor, assuming a Lambertian reflector, and $D = \frac{\varphi}{N}$ is detection system related factor. For typical values of standard grating efficiencies and numerical apertures used

in this device, the total attenuation would be approximately 60 dB. This number can be significantly improved by using custom designed diffraction gratings.

Detection will now be discussed.

Direct Spectral Measurement

The reflectance from the sample as a function of transverse location is determined by measuring the reflected spectrum from the sample arm. A simple and efficient direct measurement system 70 is shown in Fig. 4, in which an enclosure 71 houses the system 70 components.

The light returning from the imaging position carried by the fiber 14 is collimated by the objective 72 and dispersed by the grating 34. The scanning mechanism 72 is synchronized with the probe scanning mechanism or (in a compact design) the same scanning mechanism can be used to scan the probe and the detection system. This allows direct imaging using a camera 74, such as, but not limited to, a standard, inexpensive CCD camera that attaches to the detection system housing. An intensified CCD camera (ICCD) can be used when/if the signal is too weak. Electronic scanning with computer aided imaging can be implemented if a linear array is used as a detector. In this case no moving parts are necessary in the detection system 70 and higher detector sensitivity can be achieved. The detection system 70 can be either a single detector 70, a one dimensional array of detectors 70, or a two dimensional array of detectors 70. Other camera types and other detectors known to those skilled in the art are contemplated as being within the scope of the present invention.

For low light applications, the spectrum may be measured more efficiently by incorporating the device in the sample arm of an interferometer and detecting the light transmitted through a high-resolution spectrometer at the output port of the interferometer. Higher sensitivity may be achieved through the use of heterodyne detection when the light in the reference arm is modulated. The signal detected at the interferometer output will also be modulated. High signal-to-noise ratios may be then achieved by lock-in detection on the reference arm modulation frequency.

10

15

20

PCT/US01/49704 WO 02/38040

Interference Spectroscopy

5

10

15

25

30

Another method for measuring the spectrum is interference spectroscopy or Fourier transform spectroscopy. The advantages to this type of spectroscopic detection include the ability to achieve higher spectral resolutions than direct detection methods, efficient use of the returned light, inherent modulation of the reference arm by the Doppler shift of the moving mirror, the possibility of obtaining three-dimensional information, and the capability to extract both reflectance and phase data from the sample. The ability to extract phase data from the sample may allow detection of refractive index as a function of transverse position, which could give insight into the molecular composition of the sample as well as provide an additional source of image contrast other than the reflectivity of the specimen. Also, interferometric detection has the potential to allow elimination of high order multiple scattering from the signal by coherence gating. Moreover, analysis of both the phase and amplitude of the interferometric signal allows detection of group delay and dispersion. In most cases, knowledge of the group delay gives information relating the distance of the probe to the object under inspection and dispersion provides information about the shape of the object under inspection. Finally, mechanical scanning may be eliminated if points on the object, perpendicular to the spectrally encoded axis, may be separated by coherence gating.

20 Color Embodiment

In an alternative embodiment of the present invention, shown in Fig. 5, a device 100 can be constructed that uses at least two and preferably three or more separate broadband source modules 110, for example, three sources centered at red (630nm) 102, green (540nm) 104, and blue (480nm) 106 to produce color images using this technique. It is to be understood that other colors, wavelengths and number of separate sources can be selected depending on various factors, including, but not limited to, the imaging environment, imaging target, measurements to be obtained, and the like. The three energy components can be separated after reflection from the sample 30 and recombined to form an image. Each of the source/detector modules 102, 104 and 106 for the three spectral bands transmits selected wavelength light to an optical mixer/separator 108, which selectively transmits the light toward the imaging head 109 and to the imaging optics 110 for the different colors. The light reflects off

of the sample/imaging plane 112 back through the foregoing elements and is received by a color monitor 114.

One use of this embodiment is for illuminating a stained sample with one color light and detecting the reflected or retransmitted light which may be of a different wavelength or wavelengths with one or more fibers. For example, one can illuminate across the blue spectrum to detect fluorescence. This embodiment may be most practicable where the source is not broadband, but is a scanning wavelength source.

In a variation of this embodiment, an imaging device has a plurality of probes, each probe comprising an energy source, optical fiber, lens, etc., as described in the preferred embodiment. Each fiber has a distal end that is polished at an angle different from each other such that an energy source transmitted through each fiber is focused onto a distinct target site.

Multispectral Embodiment

By acquiring multiple images at different locations with the spectrally encoded probe, spectroscopic information within the bandwidth of the illuminating source may be obtained. Since each point on the sample is encoded by a different wavelength, moving the probe while acquiring images allows multiple wavelengths to be obtained from a single point on the specimen. Accumulation of these wavelengths reflected from the sample allows construction of a hyperspectral data set for each point in the image.

Multifiber Array Embodiment

In yet a further alternative embodiment of the present invention, shown in Fig. 6, an imaging apparatus 200 is provided comprising: an elongated hollow generally cylindrical body 202; a plurality of optical fibers 204 defining an array 206 disposed at least partially within the body 202 each fiber 204 having a distal end 208; a plurality of lenses 210, each lens 210 associated with a distal end 208 of each optical fiber 204 as part of said array 206, such that each lens 210 is capable of focusing energy transmitted from an energy source (not shown) through the array 206 on a distinct position on a target sample 212. Each optical fiber 204 in the array 206 has a different length such that each distal end 208 and associated lens 210 does not substantially overlap any other lens in said array 206. This embodiment also

5

10

15

20

25

incorporates a means, such as, but not limited to, mechanical, piezoelectric transducer or the like, for rotating said array about an axis.

Method

5

15

20

25

The present invention also provides a method of method for imaging, comprising: providing an endoscopic probe as described hereinabove, introducing the probe into a patient; transmitting a source energy signal to the probe such that the energy signal is directed at a sample; receiving the reflected energy from the sample; and, detecting the reflected energy.

Kits

The present invention provides a kit for performing an endoscopic procedure, comprising a probe as described hereinabove, and at least one of the following: a disinfectant, an anesthetic, and a means for introducing the probe into a patient.

The present invention also provides a kit for performing a catheterization procedure, comprising a probe as described hereinabove, and at least one of the following: a guidewire, an introducer, a syringe, an expander, and an introducer catheter.

Applications

In one embodiment, the present invention may be deployed through a needle with a gauge threshold of 20 or higher. This embodiment would allow minimally invasive access to most internal organs for the purpose of primary diagnosis, screening, or biopsy guidance. For example, areas of the spinal cord can be viewed with the present invention because the needle gauge threshold of 20 gauge or higher is met by using the endoscope of the present invention. Previously inaccessible organs such as the ovaries might be screened using the present invention deployed through a needle. Liver, pancreas, and brain biopsies could be converted to higher yield procedures by using the present invention through a needle to localize the biopsy probe to a region more likely to yield diagnostic tissue.

Certain ophthalmological surgical procedures can only be performed through small holes in the cornea or sclera, allowing access to the iris, vitreous, retina, and other internal anatomic structures within the eye. Fetal diagnosis and/or surgery may pose less of a risk to the fetus when done using the assistance of the small diameter probe

of the present invention. With various embodiments of the present invention procedures such as mammary ductoscopy, lacrimal ductoscopy, endoscopic ENT, small joint visualization, and spinal visualization can be performed.

The present invention can be used as part of a novel catheter for use in catheter-based imaging procedures. The potentially high signal-to-noise ratio of the present invention may allow imaging of the arterial wall without proximal arterial occlusion and complete vessel purging. In such an embodiment the endoscopic probe of the present invention is incorporated in one catheter lumen to provide imaging of the blood vessel or other environment into which the catheter is inserted. A second lumen can deliver therapeutics, such as, but not limited to, thrombolytic agents, plaque removing agents, antiplatelet agents, anticoagulants, vasoactive agents, combinations thereof, and the like. Alternatively, the second lumen can admit a plaque or thrombus breaking or removal device, such as, but not limited to, an ultrasonic, laser or cauterizing probe; a set of retractable teeth forming a claw for grabbing an intravascularly located body; a flushing or suction device for removing or diluting blood or other fluid which might obstruct imaging; a means for grasping a sample of material; or a cauterizing tip or the like may be employed. Alternatively, other devices, such as but not limited to, artificial a.v. fistula, other vascular access devices, and the like may be used. A catheter according to the present invention becomes possible for intravascular use only because of the small lumen size needed for the imaging probe as discussed above. Prior imaging probes were either too large or, if sufficiently small, of inadequate resolution, to be useful in a single device. With such a device, procedures previously impossible to perform efficiently now become possible.

In an alternative embodiment of the embodiment just discussed, a catheter can be constructed using a single fiber (or multiple fibers) in one lumen, whereby the fiber can be used for photodynamic therapy; i.e., imaging as well as delivering light-based therapeutic energy during a single procedure. Such energy may be magnetic, laser, ultraviolet, infrared, fluorescent, colored or other light energy. In such a device the imaging signal can be continuous or pulsed, such as alternately with the therapeutic light energy. One skilled in the art can appreciate the different permutations of pulsing, alternating, or other sequencing of imaging and therapeutic signals. Such sequencing can be controlled externally by a microprocessor or microcomputer which can be preprogrammed or manually operated by the user.

5

10

15

20

25

In a further alternative embodiment of such a catheter is a multifiber catheter, where one fiber or fibers transmit imaging light signal and another fiber or fiber transmits therapeutic light energy. Such a multifiber catheter may be desirable to have the image be where there is little absorption across the light spectrum; in other words, it is preferable to treat where there is high absorption. As such, it may be desirable to image and treat at different light wavelengths. In such a catheter where two fibers are used, the fibers can be arranged to be coaxial or side-by-side. In such a catheter the point of view of the image may be at a different point than the point of focus of the treatment light beam. If the treatment and imaging wavelengths are different, the angle of the treatment beam incident on the grating will need to be different that the angle of the imaging beam. In this case, slight adjustment of the treatment wavelength could allow direction of the treatment beam on the sample. Another alternative would be to put a dichroic (wavelength selective) beamsplitter in the distal optics of the probe that would direct the treatment beam towards the sample, while allowing the imaging light to pass through unaltered. In order to bring the angle back to the same point of focus the angle of incidence of light on the grating can be brought back. This can be achieved by polishing the end of each fiber at a different angle so that they both aim at the same target site when passed through the diffraction grating. In a further alternative a grism can be used in the distal optics. A grism is a grating placed in direct contact with a prism. By controlling the angle and the refractive index of the prism, this optical element allows for directing light in an arbitrary direction and compensates for the angular deviation of the diffracted beam from the grating. This is a very important element of the distal probe since it allows one to be able to control where one is looking (e.g., straight ahead, side view).

It is to be understood that with these embodiments using light that other electromagnetic energy wavelengths can be used with the present invention. In certain circumstances even beta, gamma or other radiation can be used in a targeted manner for treatment of cancer or other conditions while using a probed as described hereinabove in the same catheter to image the target site. Thus, such a catheter as described may be able to reduce a costly, expertise intensive surgical procedure to an outpatient one requiring less cost and expertise.

In a broad aspect of these alternative embodiments, the present invention contemplates providing one lumen for imaging and a second lumen for biopsy or

5

10

15

irrigation to all mini-endoscopy applications, not just the cardiovascular system (catheter). For instance, in mammary ductoscopy, it was found that the second lumen was necessary for irrigation, insufflation and simultaneous imaging and biopsy.

In a further alternative embodiment, imaging can be achieved through the end of the fiber or through the side of the fiber. This can be used for a dual-purpose endoscope for imaging and biopsy or treatment.

Another alternative embodiment is a combination of an imaging device as described in the preferred embodiment and a microsurgical device.

In another embodiment of the present invention the imaging probe can be used in situations where a guidewire is typically deployed. The outside of the bundle of the catheter is a wound guidewire where the total diameter is less than about 0.3 mm. Push-pull to create the images as you advance the guidewire. The probe is the tip of the guidewire to correctly position the catheter and then to advance another catheter over the guidewire.

In another embodiment of the present invention scanned wavelength can be used, possibly in the telecommunications band. Scanning the wavelength will allow scanning of the beam at the distal end at a rapid speed. This would allow simplification of the detection electronics, since only the intensity of the light will need to be detected by a single detector as opposed to measuring the spectrum of light when broad bandwidth light is used.

One can use the time dependent output of a single detector to detect light, rather than having to use interferometry or a linear array.

Conventional probes are too large to be used in pancreatic tumor visualization without causing pancreatitis. The small size of the probe of the present invention allows it to be used in pancreatic tumor visualization while minimizing the incidence of pancreatitis.

Industrial applications

The submillimeter size of the probe of the present invention also allows for new industrial applications. One application can be the textillary weaving of a submillimeter fiber into a fabric.

30

25

An alternative embodiment of the present invention provides a multiplexed array of a plurality of fibers each fiber having an associated distal optics. More area can be scanned and the number of resolvable elements increases with this embodiment. Or, a single fiber with multiple diffraction elements spaced along the length of the fiber, e.g., a fiber grating can be used at 1 cm, a second fiber grating can be used at 2 cm, etc.

Another application is probe used as an inspection system, which comprises a cylindrical or other shape (regular or irregular) body having at least one and preferably a plurality of imaging fibers extending axially outward from the body in a regular or irregular array or arrays. Alternatively, the fibers can be flush or minimally protruding from the body. At one end of the body is an opening through which either the fibers or fiber passes which is connected to the detection apparatus. This detector fiber array can be used in pipes, conduits, or other closed or open systems not previously accessible to image longitudinally, three-dimensionally, panoramically, stereoscopically, and the like, using the multiple fiber array to image multiple points. This would allow for much more surface area and volume to be imaged and analyzed in a single procedure than previously possible. An array of this type can analyze inner wall defects in conduits and the like.

In a variation on this embodiment, shown in Fig. 7, a probe has a single optical fiber 300 which has a plurality of grating lenses 302(or other lenses) spaced along the surface 304 of the fiber 300. Each lens 302a, b, c, d, n focuses energy onto a distinct target site 306a, b, c, d, n.

Fluid properties can be measured dynamically using the present invention. Such an application can have a multifiber array probe within a very small tube. A dye or other detectable substance can be passed through the tube and the probe illuminate the fluid and detectable substance to obtain fluid flow dynamics in a given environment, such as were a weakness in the wall or partial obstruction has occurred. The present invention can be adapted to provide a surface built into the probe which can reflect illuminated light which has passed through an aliquot of fluid, thus permitting absorption measurements to be taken. Turbidity, color change, and other conditions may be detected in situ in small vessels, for example, kidney and gall bladder conditions (e.g., stones), seminal fluid composition, and the like. Alternatively, a cell can be constructed which can continuously admit and pass a fixed volume of fluid,

5

10

15

20

25

thus permitting accurate measurement to be made of the cell volume in situ without requiring surgery, disruption or occlusion of the vessel. Much more accurate diagnosis can be made with the present invention rather than external measurements as must currently be made. With the small size of the present invention, coupled with the high resolution attainable, spaces previously impractical to be imaged can now be visualized.

The present invention can be developed into a self-contained remote controlled imaging probe where image data is transmitted by radio frequency or other signal from within the tube or vessel to an external receiver.

The present invention can also be used in veterinary applications for performing endoscopic procedures on small animals, fish, birds, reptiles and the like.

Advantages

5

15

20

The present invention provides for an imaging device capable of imaging at video rates or higher with up to or exceeding about one hundred times the number of resolvable points of currently available fiber optic bundles. The probe of the present invention is single mode fiber based and can be flexible or rigid, according to required working parameters. No fill factor problem is encountered because a single fiber is used. Dead pixels are eliminated in the present invention. Light transmission efficiency is maintained because there is no crosstalk compared to multiple fiber bundles. High efficiency imaging, allowing clear visualization through turbid fluid may be obtained through heterodyne detection. Analysis of the group delay and dispersion of the light returned from the sample allows the acquisition of three-dimensional information from the object. Hyperspectral data may be obtained from a series of offset images.

- The present invention can be used to convert many current surgical procedures into outpatient based procedures. Biopsy guidance, cancer screening, e.g., abdominal cavity, intracranial, spinal cord, are possible. The present invention is able to reach into spaces too small for current endoscopes and maintain a useful resolution level of obtainable information.
- The probe of the present invention can fit through a 20 or higher gauge needle. Due to the small size of the probe, it may also not require anesthesia. Physicians and other

medical personnel may be able perform procedures not previously doable by endoscopy. Also, lesser trained personnel may now be able to perform procedures previously performable only by specially trained physicians and/or surgeons. With the present invention one can package an endoscope having superior resolution in a diameter smaller than 1 mm, yet reduce the number of physical components that make up the endoscope.

Although only a few exemplary embodiments of this invention have been described in detail above, those skilled in the art will readily appreciate that many modifications are possible in the exemplary embodiments without materially departing from the novel teachings and advantages of this invention. Accordingly, all such modifications are intended to be included within the scope of this invention as defined in the following claims. All patents, applications, publications and other documents referred to herein are incorporated by reference in their entirety.

5

CLAIMS

Claimed is:

1. A spectrally encoded endoscopic probe capable of having spatially encoded location information, comprising:

- 5 a) at least one flexible energy conducting member;
 - b) a source of energy;
 - a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
- d) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis;
 - e) means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
- 15 f) means for receiving energy reflected from said sample.
 - 2. The probe of Claim 1, wherein said at least one flexible energy conducting member comprises at least one optical fiber.
 - 3. The probe of Claim 1, wherein said fiber is mode or phase modulated.
- 4. The probe of Claim 1, wherein said source of energy is a light-emitting diode, super-luminescent diode, rare-earth doped fibers, solid-state mode-locked laser, spectrally broadened laser, monochromatic light, polychromatic light, infrared, ultraviolet, ultrasonic, low or high energy radiation, x-ray radiation, alpha radiation, beta radiation, or gamma radiation, or mixtures thereof.
 - 5. The probe of Claim 1, wherein said dispersive element is a diffractive element.
- 25 6. The probe of Claim 1, wherein said dispersive element is a refractive element.

7. The probe of Claim 1, wherein said dispersive element is a fiber grating, blazed grating, binary, prism, grism or holographic lens grating.

- 8. The probe of Claim 1, wherein said means for focusing comprises a lens.
- The probe of Claim 1, wherein said lens is a gradient index lens, a reflective
 mirror lens grating combination or diffractive lens.
 - 10. The probe of Claim 1, wherein said means for scanning is a piezoelectric transducer or a torque transducing device.
 - 11. The probe of Claim 1, further comprising means for detecting said received reflected energy.
- 10 12. The probe of Claim 11, wherein said detection means is a single detector, one dimensional array of detectors or a two dimensional array of detectors.
 - 13. The probe of Claim 12, wherein said detection means is a means for interferometric spectral decoding.
- 14. The probe of Claim 12, wherein said detection means is a means for direct spectral decoding.
 - 15. The probe of Claim 1, further comprising a mirror.
 - 16. The probe of Claim 1, further comprising a means for polarization control.
 - 17. The probe of Claim 1, further comprising a beam splitter.
 - 18. The probe of Claim 1, further comprising a beam stop.
 - 20 19. The probe of Claim 11, wherein said detection means is physically associated with said probe.
 - 20. The probe of Claim 11, wherein said detection means provides spectroscopic information.
 - 21. The probe of Claim 11, wherein said detection means provides three dimensional information.

22. The probe of Claim 1, wherein said probe has a diameter of less than about 1.0 mm.

- 23. The probe of Claim 1, wherein said probe has a number of resolvable points of from about 10,000 to about 1,000,000.
- 5 24. The probe of Claim 1, wherein said probe has a number of resolvable points of from about 150,000 to about 300,000.
 - 25. The probe of Claim 1, wherein said probe has a number of resolvable points of from about 10,000 to about 150,000.
- 26. A spectrally encoded endoscopic probe capable of having spatially encoded location information, comprising:
 - a) a body having a proximal end and a distal end;
 - b) an elongated flexible energy conducting member having a proximal end and a distal end;
- c) an optical head associated with said distal end of said energy conducting member, said optical head being capable of rotatable or translational movement with respect to said body.

- 27. A method for imaging, comprising:
 - a) providing an endoscopic probe capable of having spatially encoded location information, comprising:
 - i) at least one flexible energy conducting member;
- 5 ii) a source of energy;
 - iii) a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
 - iv) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis;
 - means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
 - vi) means for receiving energy reflected from said sample;
- b) introducing said probe into a patient;
 - c) transmitting a source energy signal to said probe such that said energy signal is directed at a sample;
 - d) receiving the reflected energy from said sample; and,
 - e) detecting said reflected energy.
- 20 28. The method of Claim 27, wherein said probe has a diameter of less than about 1.0 mm.
 - 29. The method of Claim 27, wherein said probe has a number of resolvable points of from about 300,000 to about 1,000,000.
- The method of Claim 27, wherein said probe has a number of resolvable points of from about 150,000 to about 300,000.

31. The method of Claim 27, wherein said probe has a number of resolvable points of from about 100,000 to about 150,000.

- 32. A detection system using spectrally encoded information, comprising:
 - a) a flexible light conducting member;
- 5 b) a housing;
 - c) means for focusing energy;
 - d) means for dispersing energy received from said means for focusing energy; and,
 - e) means for scanning.

33. An imaging device capable of detecting a plurality of wavelengths of energy reflected from a sample, comprising:

- a plurality of probes, each probe capable of having spatially encoded location information and comprising at least one flexible energy conducting member;
- ii) a source of energy;
- iii) a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
- iv) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis;
- v) means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
- vi) means for receiving energy reflected from said sample;
- b) a plurality of wavelengths of energy capable of impinging on said sample;
- c) wherein each energy delivering fiber has an end that is polished at an angle different from each other such that an energy source transmitted through each fiber is focused onto a single target site.

5

10

15

34. An imaging device capable of detecting a plurality of wavelengths of energy reflected from a sample, comprising:

- a) an endoscopic probe capable of having spatially encoded location information, comprising:
 - i) at least one flexible energy conducting member;
 - ii) a source of energy;
 - iii) a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
 - iv) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis;
 - v) means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
 - vi) means for receiving energy reflected from said sample;
- b) at least one energy source capable of producing a plurality of wavelengths of energy capable of impinging on said sample; and,
- c) a plurality of focusing means associated with and spaced along said fiber such that each focusing means is capable of focusing energy on a distinct location on said sample.

5

10

- 35. A probe, comprising:
 - a) at least one lumen;
 - b) a spectrally encoded imaging probe comprising
 - i) at least one flexible energy conducting member;
- 5 ii) a source of energy;
 - iii) a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
 - iv) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis;
 - v) means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
 - vi) means for receiving energy reflected from said sample; and,
- 15 c) means for introducing said catheter through the skin and into a blood vessel of a patient.
 - 36. The probe of Claim 35, wherein said at least one lumen comprises a first lumen and a second lumen, said first lumen capable of containing said probe, said second lumen capable of delivering an agent or device to a target area.
- 20 37. The probe of Claim 23, wherein said catheter has a diameter of less than or equal to about 1.0 mm.
 - 38. The probe of Claim 36, wherein said probe has a diameter of less than about 1.0 mm.
- The probe of Claim 35, wherein said probe has a resolution of from about 300,000 to about 1,000,000 resolvable points.

40. The probe of Claim 35, wherein said probe has a resolution of from about 150,000 to about 300,000 resolvable points.

- 41. The probe of Claim 35, wherein said probe has a resolution of from about 100,000 to about 150,000 resolvable points.
- 5 42. The probe of Claim 35, wherein said agent is a drug.
 - 43. The probe of Claim 42, wherein said drug is a thrombolytic agent, plaque removing agent, antiplatelet agent, anticoagulant, vasoactive agent, or a combination thereof.
 - 44. The probe of Claim 35, wherein said agent is a device.
- 10 45. The probe of Claim 44, wherein said device is an ultrasonic, laser or cauterizing probe, a set of retractable teeth forming a claw for grabbing an intravascularly located body, a suction tube, a means for grasping a sample of material, a cauterizing tip or an artificial a.v. fistula.
- 46. The probe of Claim 35, wherein said agent is energy provided by an energy source.
 - 47. The probe of Claim 35, further comprising means for displacing fluid from the field of view.
 - 48. A multifiber catheter having at least one imaging fiber and at least one therapeutic light energy delivering fiber.
- 20 49. The multifiber catheter of Claim 48, wherein said imaging fiber is capable of transmitting energy at a first wavelength and said therapeutic light energy delivering fiber is capable of transmitting energy at a second wavelength.
 - 50. The multifiber catheter of Claim 48, wherein said imaging fiber and said energy delivering fiber are coaxial.
- 25 51. The multifiber catheter of Claim 48, wherein said imaging fiber and said energy delivering fiber are in a side-by-side configuration.

PCT/US01/49704 WO 02/38040

52. The multifiber catheter of Claim 48, wherein said imaging fiber and said energy delivering fiber each have an end that is polished at an angle different from each other such that an energy source passing through each fiber is focused onto a single target site.

- 5 53. The catheter of Claim 48, wherein said first wavelength and said second wavelength are different.
 - 54. The catheter of Claim 48, wherein said first wavelength and said second wavelength are the same.
- 55. A multifiber imaging apparatus using spectrally encoded information,comprising:
 - a) an elongated hollow generally cylindrical body having a plurality of spaced apart apertures defined on the surface thereon;
 - b) a plurality of flexible energy conducting fibers disposed at least partially within said body, at least one fiber positioned at least partially within each of said apertures;
 - c) an imaging head associated with each of said fibers; and,
 - d) at least one detector associated with said plurality of fibers.
 - 56. An imaging apparatus, comprising:
 - a) an elongated hollow generally cylindrical body;
 - 20 b) a plurality of optical fibers defining an array disposed at least partially within said body each fiber having a distal end;
 - a plurality of lenses, each lens associated with a distal end of each optical fiber as part of said array,
 - such that each lens is capable of focusing energy transmitted from an energy source through said array on a distinct position on a target sample.

57. The imaging apparatus of Claim 56, wherein each optical fiber in said array has a different length such that each distal end and associated lens does not substantially overlap any other lens in said array.

- 58. The imaging apparatus of Claim 56, further comprising means for rotating said array about an axis.
- 59. An imaging apparatus, comprising:
 - a) an optical fiber having an outer surface; and,
 - b) a plurality of means for focusing a source of energy onto a distinct target position; each focusing means being spaced along said outer surface, wherein said energy source is spectrally encoded.

33

5

60. A remote controlled spectrally encoded imaging system, comprising:

- a) an endoscopic probe capable of having spatially encoded location information, comprising:
 - i) at least one flexible energy conducting member;

5 ii) a source of energy;

- iii) a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
- iv) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis;
 - v) means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
 - vi) means for receiving energy reflected from said sample;
- b) means for detecting the image information from said transmitted information.
 - means associated with said probe for transmitting the detected information;
 - means for receiving information transmitted by said probe; and,
- 20 e) means for processing said information.
 - 61. The imaging system of Claim 60, wherein said remote controlled spectrally detection means provides spectroscopic information.
 - 62. The imaging system of Claim 60, detection means provides three dimensional information.

63. The imaging system of Claim 60, wherein said detection means is a single detector, one dimensional array of detectors or a two dimensional array of detectors.

- 64. The imaging system of Claim 60, wherein said detection means is a means for interferometric spectral decoding.
 - 65. The imaging system of Claim 60, wherein said detection means is a means for direct spectral decoding.

10

- 66. A kit for performing an endoscopic procedure, comprising:
 - an endoscopic probe capable of having spatially encoded location information, comprising:
 - i) at least one flexible energy conducting member;
- 5 ii) a source of energy;
 - iii) a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
 - iv) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis; and,
 - v) means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
 - b) means for receiving energy reflected from said sample;
 - c) a disinfectant;
 - d) an anesthetic; and,
 - e) means for introducing said probe into a patient.

67. A kit for performing a catheterization procedure, comprising:

a) an endoscopic probe capable of having spatially encoded location information, comprising:

- i) at least one flexible energy conducting member;
- ii) a source of energy;
 - iii) a dispersive element through which said energy is transmitted or reflected such that said energy spectrum is dispersed;
 - iv) means for focusing said dispersed energy onto a sample such that the impingement spot for each wavelength is at a distinct location on said sample, the spectrum of wavelength defining a wavelength encoded axis;
 - v) means for scanning said sample with said focused energy in a direction different from said wavelength encoded axis; and,
 - vi) means for receiving energy reflected from said sample;
- b) a guidewire;
 - c) an introducer;
 - d) a syringe;
 - e) at least one expander; and,
 - f) an introducer catheter.

5

10

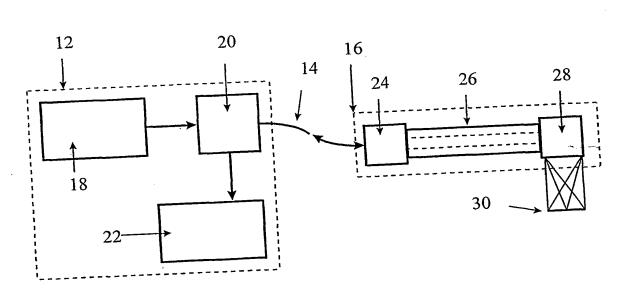


FIG. 1

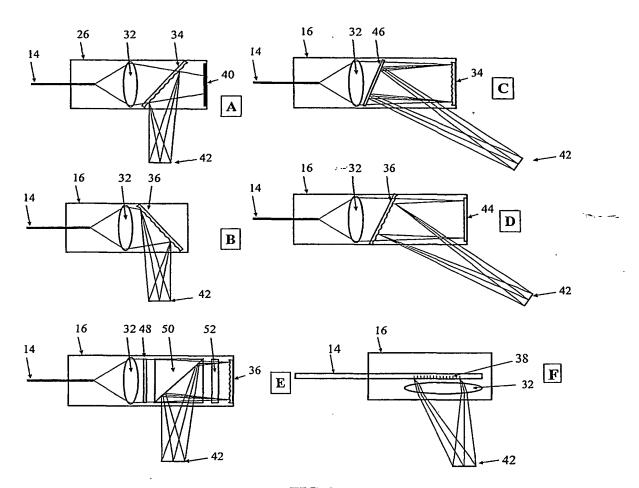


FIG. 2

PCT/US01/49704 WO 02/38040

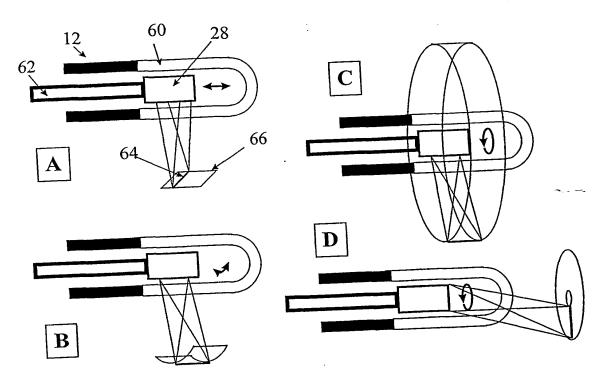
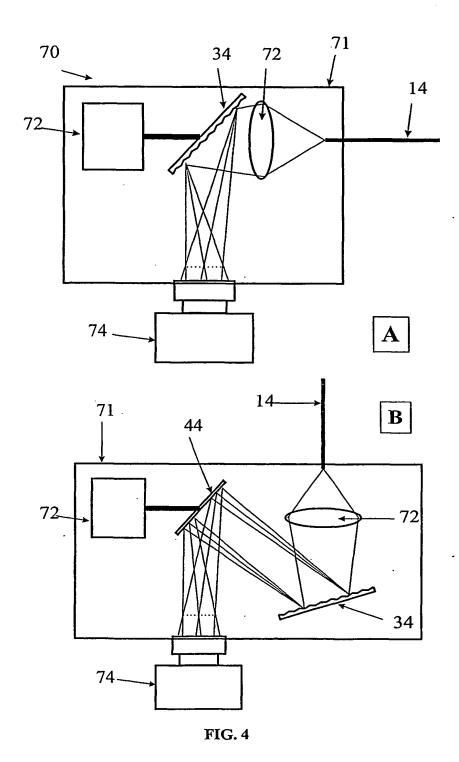


FIG. 3



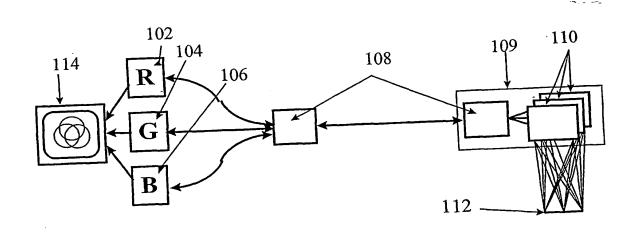


FIG. 5

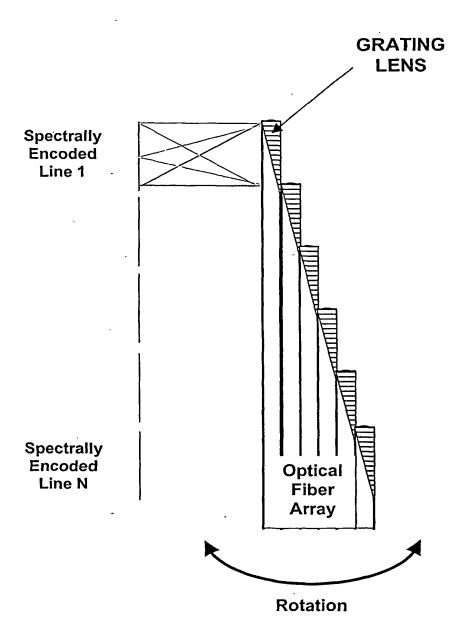


FIG. 6

PCT/US01/49704

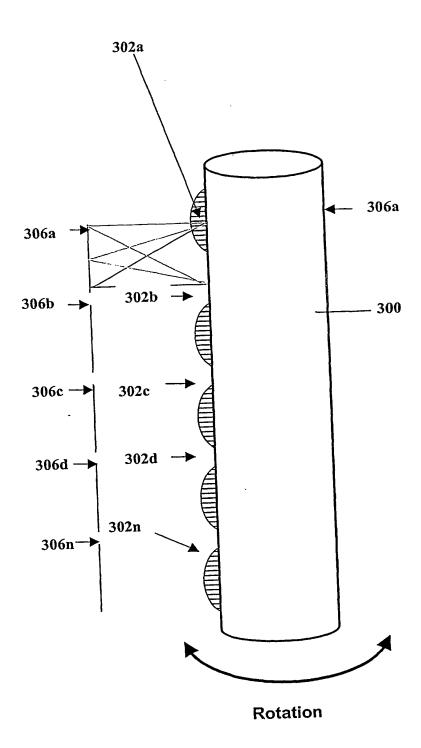


FIG. 7

(19) World Intellectual Property Organization International Bureau





(43) International Publication Date 16 May 2002 (16.05.2002)

PCT

(10) International Publication Number WO 02/038040 A3

(51) International Patent Classification⁷: A61B 1/313, 1/07, 1/018, 5/00, G02B 21/00, 23/24

(21) International Application Number: PCT/US01/49704

(22) International Filing Date:

9 November 2001 (09.11.2001)

(25) Filing Language:

English

(26) Publication Language:

English

(30) Priority Data: 09/709,162 10 November 2000 (10.11.2000) US

(71) Applicant (for all designated States except US): THE GENERAL HOSPITAL CORPORATION [US/US]; 55 Fruit Street, Boston, MA 02114 (US).

(72) Inventors; and

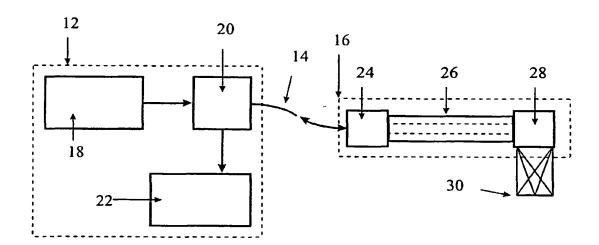
(75) Inventors/Applicants (for US only): TEARNEY, Guillermo, J. [US/US]; 118 Kinnaird Steet Number 3, Cambridge, MA 02139 (US). BOUMA, Brett, Eugene [US/US]; 12 Monmouth Street, Quincy, MA 02171 (US). SHISKOV, Milen, Stefanov [BG/US]; 122 Bowdoin Street, Apt. 2, Boston, MA 02108 (US). ROSEN, Jonathan, Jay [US/US]; 12 Surrey Road, Newton, MA 02458 (US).

(74) Agent: BERNSTEIN, Jason, A.; Bernstein & Associates, P.C., Suite 495, Embassy Row 400, 6600 Peachtree Dunwoody Road, N.E., Atlanta, GA 30328-1649 (US).

- (81) Designated States (national): AE, AG, AL, AM, AT (utility model), AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ (utility model), CZ, DE (utility model), DE, DK (utility model), DK, DM, DZ, EC, EE (utility model), EE, ES, FI (utility model), FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG; KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK (utility model), SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.
- (84) Designated States (regional): ARIPO patent (GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW), Eurasian patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), European

[Continued on next page]

(54) Title: SPECTRALLY ENCODED MINIATURE ENDOSCOPIC IMAGING PROBE



(57) Abstract: A spectrally encoded endoscopic probe having high resolution and small diameter comprising at least one flexible optical fiber; an energy source; a grating through which said energy is transmitted such that the energy spectrum is dispersed; a lens for focusing the dispersed energy spectrum onto a sample such that the impingement spot for each wavelength is a separate position on the sample, the wavelength spectrum defining a wavelength encoded axis; means for mechanically scanning the sample with focused energy in a direction perpendicular to the wavelength encoded axis; a means for receiving energy reflected from the sample; and, a means for detecting the received reflected energy. The probe grating and lens delivers a beam of multispectral light having spectral components extending in one dimension across a target region and which is moved to scan in another direction. The reflected spectrum is measured to provide two dimensional imaging of the region.

O 02/038040 A



patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR), OAPI patent (BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG).

Published:

— with international search report

(88) Date of publication of the international search report: 27 February 2003

For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

INTERNATIONAL SEARCH REPORT

onal Application No PCT/US 01/49704

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61B1/313 A61B1/07 G02B23/24

A61B1/018

A61B5/00

G02B21/00

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

 $\begin{tabular}{ll} Minimum documentation searched (classification system followed by classification symbols) \\ IPC 7 & A61B & G02B & F21V \end{tabular}$

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

EPO-Internal, PAJ, WPI Data

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X Y	WO 99 44089 A (GEN HOSPITAL CORP ;TEARNEY GUILLERMO J (US); BOUMA BRETT E (US); W) 2 September 1999 (1999-09-02) cited in the application the whole document	1-6,8, 10-15, 17-26, 32,60-65 35-47, 66,67
x	US 5 785 651 A (BAKER PHILLIP C ET AL) 28 July 1998 (1998-07-28)	1-9,11, 12, 14-17, 19-21, 32,33,
	column 6, line 27 -column 7, line 3 column 7, line 39 -column 7, line 59 column 10, line 1 -column 10, line 30; claims 5-8,12; figures 3,5,9,10	60-63,65
	 -/	

X Further documents are listed in the continuation of box C.	X Patent family members are listed in annex.
 Special categories of cited documents: "A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed 	"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. "&" document member of the same patent family
Date of the actual completion of the international search 29 August 2002	Date of mailing of the international search report 1 0. 12. 02
Name and mailing address of the ISA European Patent Office, P.B. 5818 Patentlaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Authorized officer Rick, K

Form PCT/ISA/210 (second sheet) (July 1992)

INTERNATIONAL SEARCH REPORT

Int onal Application No . PCT/US 01/49704

	INTERNATIONAL SEARCH REPORT	PCT/US 01/49704 -		
10 - stimus	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	Relevant to claim No.		
ategory °	Citation of document, with indication, where appropriate, of the relevant passages	•		
X	US 4 585 349 A (GROSS DANIEL ET AL) 29 April 1986 (1986-04-29)	1-9,11, 12, 14-17, 19-21, 32, 60-63,65		
	column 6, line 28 -column 7, line 7; claims 4-14; figure 3	35-47,		
Y A	US 5 817 144 A (GREGORY KENTON W) 6 October 1998 (1998-10-06) column 4, line 66 -column 5, line 45 column 6, line 61 -column 7, line 35;	66,67 34,59		
	Claim 12; Figures 1	35-41, 44-47,		
Y	DE 195 42 955 A (POLYDIAGNOST VERTRIESS & SERVI; SCHWIND GMBH & CO KG HERBERT (DE)) 22 May 1997 (1997-05-22) 22 May 1997 (1997-05-22) 23 column 1, line 58 -column 3, line 61	66,67 34,59		

Form PCT/ISA/210 (continuation of second sheet) (July 1992)

national application No. PCT/US 01/49704

INTERNATIONAL SEARCH REPORT

Observations where certain claims were found unsearchable (Continuation of item ◆ of first sheet)					
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:					
1. X Claims Nos.: 27-31 because they relate to subject matter not required to be searched by this Authority, namely: Rule 39.1(iv) PCT - Method for treatment of the human or animal body by					
2. Claims Nos.: because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:					
3. Claims Nos.: because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).					
Box II Observations where unity of invention is lacking (Continuation of item 2 of first sheet)					
This International Searching Authority found multiple inventions in this international application, as follows:					
see additional sheet					
As all required additional search fees were timely paid by the applicant, this International Search Report covers all searchable claims.					
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.					
3. As only some of the required additional search fees were timely paid by the applicant, this International Search Report covers only those claims for which fees were paid, specifically claims Nos.:					
4. No required additional search fees were timely paid by the applicant. Consequently, this International Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.: 1-26,32-47,59-67					
Remark on Protest The additional search fees were accompanied by the applicant's protest. No protest accompanied the payment of additional search fees.					

Form PCT/ISA/210 (continuation of first sheet (1)) (July 1998)

FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. Claims: 1-26,32-47,59-67

A spectrally encoded endoscopic probe.

2. Claims: 48-54

A multifiber catheter having at least one imaging and one therapeutic light energy delivering fiber.

3. Claim: 55

A multifiber catheter having a plurality of spaced apart apertures on the surface of an elongated hollow cylindrical body.

4. Claims: 56-58

A multifiber catheter having an array of optical fibers and lenses associated with the distal end of each fiber.

BNSDOCID <WO____0238040A3_I_>

INTERNATIONAL SEARCH REPORT

Information on patent family members

Intension No
PCT/US 01/49704

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 9944089	A	02-09-1999	AU EP JP WO US	2882399 A 1057063 A1 2002505434 T 9944089 A1 6341036 B1 2002122246 A1	15-09-1999 06-12-2000 19-02-2002 02-09-1999 22-01-2002 05-09-2002
US 5785651	A	28-07-1998	AU EP WO	5860296 A 0830565 A1 9641123 A1	30-12-1996 25-03-1998 19-12-1996
US 4585349	Α	29-04-1986	CH EP JP	663466 A5 0142464 A1 60073405 A	15-12-1987 22-05-1985 25-04-1985
US 5817144	Α	06-10-1998	US US	5571151 A 5836940 A	05-11-1996 17-11-1998
DE 19542955	Α	22-05-1997	DE	19542955 A1	22-05-1997

Form PCT/ISA/210 (patent lamily annex) (July 1992)

THIS PAGE LEFT BLANK